



Robert Greene Sterne
Edward J. Kessler
Jorge A. Goldstein
David K.S. Cornwell
Robert W. Esmond
Tracy-Gene G. Durkin
Michele A. Cimbala
Michael B. Ray
Robert E. Sokohl
Eric K. Steffe
Michael Q. Lee
Steven R. Ludwig
John M. Covert
Linda E. Alcorn
Robert C. Milonig
Lawrence B. Bugaisky
Donald J. Featherstone
Michael V. Messinger

Judith U. Kim
Timothy J. Shea, Jr.
Patrick E. Garrett
Jeffrey T. Helvey*
Heidi L. Kraus
Edward W. Yee
Albert L. Ferro*
Donald R. Banowitz
Peter A. Jackman
Molly A. McCall
Teresa U. Medler
Jeffrey S. Weaver
Kendrick P. Patterson
Vincent L. Capuano
Albert J. Fasulo II*
Eldora Ellison Floyd
Thomas C. Fiala
Brian J. Del Buono

Virgil Lee Beaston*
Kimberly N. Reddick
Theodore A. Wood
Elizabeth J. Haanes
Bruce E. Chalker
Joseph S. Ostroff
Frank R. Cottingham
Christine M. Lhulier
Rae Lynn Prengaman
Jane Shershenovich*
Lawrence J. Carroll*
George S. Bardmesser
Daniel A. Klein*
Rodney G. Maze
Jason D. Eisenberg
Michael A. Specht*
Andrea J. Kamage
Tracy L. Muller

Jon E. Wright*
LuAnne M. Yuricek*
Registered Patent Agents*
Karen R. Markowicz
Nancy J. Leith
Ann E. Summerfield
Helene C. Carlson
Gaby L. Longworth
Matthew J. Dowd
Aaron L. Schwartz
Angelique G. Uy
Boris A. Matvenko
Mary B. Tung
Katrina Y. Pei
Bryan L. Skelton
Robert A. Schwartzman
John J. Figueroa

Timothy A. Doyle
Jennifer R. Mahalingappa
Teresa A. Colella
Jeffrey S. Lundgren
Victoria S. Rutherford
W. Brian Edge

Senior Counsel
Samuel L. Fox
Kenneth C. Bass III
Lisa A. Dunner

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*Admitted only in Virginia
*Admitted only in Texas
*Practice Limited to
Federal Agencies

March 5, 2003

WRITER'S DIRECT NUMBER:
(202) 772-8560

INTERNET ADDRESS:
RESMOND@SKGF.COM

Commissioner for Patents
Washington, D.C. 20231

Hand Carry to Examiner W. Moore
Art Unit 1652

Re: U.S. Utility Patent Application
Appl. No. 10/024,149; Filed: December 21, 2001
For: **Cloned Genes Encoding Reverse Transcriptase
Lacking RNase H Activity**
Inventors: KOTEWICZ *et al.*
Our Ref: 0942.049000A/RWE/MTT

Sir:

Transmitted herewith for appropriate action are the following documents:

1. Information Disclosure Statement and list of cited references on Form PTO-1449 (13 sheets);
2. First Supplemental Information Disclosure Statement and list of cited references on Form PTO-1449 (3 sheets);
3. Second Supplemental Information Disclosure Statement and list of cited references on Form PTO-1449 (11 sheets);
4. Third Supplemental Information Disclosure Statement and list of cited references on Form PTO-1449 (3 sheets);
5. Fourth Supplemental Information Disclosure Statement and list of cited references on Form PTO-1449 (26 sheets);
6. Fifth Supplemental Information Disclosure Statement and list of cited references on Form PTO-1449 (1 sheet);

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March 5, 2003

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7. Sixth Supplemental Information Disclosure Statement and list of cited references on Form PTO-1449 (15 sheets);
8. Seventh Supplemental Information Disclosure Statement and list of cited references on Form PTO-1449 (3 sheets);
9. Eighth Supplemental Information Disclosure Statement and list of cited references on Form PTO-1449 (32 sheets);
10. Ninth Supplemental Information Disclosure Statement and list of cited references on Form PTO-1449 (8 sheets);
11. Tenth Supplemental Information Disclosure Statement;
12. Eleventh Supplemental Information Disclosure Statement and list of cited references on Form PTO-1449 (2 sheets);
13. Twelfth Supplemental Information Disclosure Statement and list of cited references on Form PTO-1449 (1 sheet);
14. Thirteenth Supplemental Information Disclosure Statement and list of cited references on Form PTO-1449 (5 sheets);
15. Fourteenth Supplemental Information Disclosure Statement, list of cited references on Form PTO-1449 (5 sheets), and a copy of each reference cited on Form PTO-1449 (21 documents: AH1-AB2, AM1, and AS112-AT116); and
16. One (1) return postcard.

It is respectfully requested that the attached postcard be stamped with the date of filing of these documents, and that it be returned to our courier. In the event that extensions of time are necessary to prevent abandonment of this patent application, then such extensions of time are hereby petitioned.

Commissioner for Patents

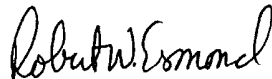
March 5, 2003

Page 3

The U.S. Patent and Trademark Office is hereby authorized to charge any fee deficiency, or credit any overpayment, to our Deposit Account No. 19-0036.

Respectfully submitted,

STERNE, KESSLER, GOLDSTEIN & FOX P.L.L.C.

A handwritten signature in black ink, appearing to read "Robert W. Esmond". The signature is fluid and cursive, with the first name "Robert" and last name "Esmond" clearly distinguishable.

Robert W. Esmond
Attorney for Applicants
Registration No. 32,893

RWE/MTT/acr
Enclosures

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Dmt
H-10-93

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re application of:

Kotewicz *et al.*

Appl. No. 10/024,149

Filed: December 21, 2001

For: **Cloned Genes Encoding Reverse
Transcriptase Lacking RNase H
Activity**

Confirmation No. 4033

Art Unit: 1652

Examiner: *To Be Assigned*

Atty. Docket: 0942.049000A/RWE/MTT

Information Disclosure Statement

Assistant Commissioner for Patents
Washington, D.C. 20231

Sir:

Listed on accompanying Form PTO-1449 are documents that may be considered material to the examination of this application, in compliance with the duty of disclosure requirements of 37 C.F.R. §§ 1.56, 1.97 and 1.98. All documents were cited by or submitted to the Office in Application No. 08/798,458, filed February 10, 1997, which issued as U.S. Patent No. 6,063,608; Application No. 08/614,260, filed March 12, 1996, which issued as U.S. Patent No. 5,668,005, on September 16, 1997; Application No. 08/404,907, filed March 15, 1995; Application No. 07/671,156, filed March 18, 1991, which issued as U.S. Patent No. 5,244,797 on September 14, 1993, and was reexamined as Control No. 90/003,803; and Application No. 07/825,260, filed January 24, 1992, which issued as U.S. Patent No. 5,405,776 on April 11, 1995 and was reexamined as Control No. 90/003,804. Copies of the cited documents may be found in these parent applications and are not enclosed.

Where the publication date of a listed document does not provide a month of publication, the year of publication of the listed document is sufficiently earlier than the effective U.S. filing date and any foreign priority date so that the month of publication is not in issue. Applicants have listed publication dates on the attached PTO-1449 based on information presently available to the undersigned. However, the listed publication dates should not be construed as an admission that the information was actually published on the date indicated.

Applicants reserve the right to establish the patentability of the claimed invention over any of the information provided herewith, and/or to prove that this information may not be prior

art, and/or to prove that this information may not be enabling for the teachings purportedly offered.

This statement should not be construed as a representation that a search has been made, or that information more material to the examination of the present patent application does not exist. The Examiner is specifically requested not to rely solely on the material submitted herewith. It is further understood that the Examiner will consider information that had been cited or submitted to the U.S. Patent and Trademark Office in a prior application relied on under 35 U.S.C. § 120. 1138 OG 37, 38 (May 19, 1992).

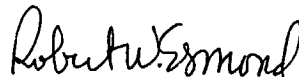
This Information Disclosure Statement is being filed more than three months after the U.S. filing date but before the mailing date of a first Office Action on the merits. No statement or fee is required.

It is respectfully requested that the Examiner initial and return a copy of the enclosed PTO-1449, and to indicate in the official file wrapper of this patent application that the documents have been considered.

The U.S. Patent and Trademark Office is hereby authorized to charge any fee deficiency, or credit any overpayment, to our Deposit Account No. 19-0036.

Respectfully submitted,

STERNE, KESSLER, GOLDSTEIN & FOX P.L.L.C.



Robert W. Esmond
Attorney for Applicants
Registration No. 32,893

Date: March 5, 2003

1100 New York Avenue, N.W.
Suite 600
Washington, D.C. 20005-3934
(202) 371-2600

FORM PTO-1449 <u>INFORMATION DISCLOSURE STATEMENT</u>	ATTY. DOCKET NO. 0942.049000A/RWE/MTT	APPLICATION NO. 10/024,149
	APPLICANTS Kotewicz et al.	
	FILING DATE December 21, 2001	GROUP 1652

U.S. PATENT DOCUMENTS

EXAMINER INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUB- CLASS	FILING DATE
	AA1	4,663,290	05/05/1987	Weis et al.	435	253	01/21/1982
	AB1	4,795,699	01/03/1989	Tabor et al.	435	5	01/14/1987
	AC1	4,943,531	07/24/1990	Goff et al.	435	194	05/06/1985
	AD1	5,017,492	05/21/1991	Kotewicz et al.	435	252.3	02/27/1986
	AE1	5,244,797	09/14/1993	Kotewicz et al.	435	194	03/18/1991
	AF1	5,405,776	04/11/1995	Kotewicz et al.	435	252.33	01/24/1992
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FOREIGN PATENT DOCUMENTS

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	AL						Yes No
	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>1</u>	Akins, R.A. et al., "Mitochondrial Plasmids of Neurospora: Integration into Mitochondrial DNA and Evidence for Reverse Transcription in Mitochondria," Cell 47:505-516, Cell Press (1986).
	AS	<u>1</u>	Bassin, R.H. et al., "Macromolecular Requirements for Abrogation of Fv-1 Restriction by Murine Leukemia Viruses," J. Virology 35(2):287-297, American Society for Microbiology (1980).
	AT	<u>1</u>	Berger, S.L. et al., "Reverse Transcriptase and Its Associated Ribonuclease H: Interplay of Two Enzyme Activities Controls the Yield of Single-Stranded Complementary Deoxyribonucleic Acid," Biochemistry 23:2365-2372, American Chemical Society (1983).

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	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>2</u>	Blain, S.W. and Goff, S.P., "Nuclease Activities of Moloney Murine Leukemia Virus Reverse Transcriptase," <i>J. Biol. Chem.</i> 268(31):23585-23592, American Society for Biochemistry and Molecular Biology, Inc. (1993).
	AS	<u>2</u>	Carter, P. and Wells, J.A., "Engineering Enzyme Specificity by 'Substrate-Assisted Catalysis,'" <i>Science</i> 237:394-399, American Association for the Advancement of Science (1987).
	AT	<u>2</u>	Fawcett, D.H. et al., "Transposable Elements Controlling I-R Hybrid Dysgenesis in <i>D. melanogaster</i> Are Similar to Mammalian LINES," <i>Cell</i> 47:1007-1015, Cell Press (1986).

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	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>3</u>	Finston, W.I. and Champoux, J.J., "RNA-Primed Initiation of Moloney Murine Leukemia Virus Plus Strands by Reverse Transcriptase In Vitro," <i>J. Virology</i> 51(1):26-33, American Society for Microbiology (1984).
	AS	<u>3</u>	Gerard, G.F. et al., "Poly (2'-O-methylcytidylate) Oligodeoxyguanylate as a Template for the Ribonucleic Acid Directed Deoxyribonucleic Acid Polymerase in Ribonucleic Acid Tumor Virus Particles and a Specific Probe for the Ribonucleic Acid Directed Enzyme in Transformed Murine Cells," <i>Biochemistry</i> 13(8):1632-1641, Academic Press, Inc. (1974).
	AT	<u>3</u>	Gerard, G.F. and Grandgenett, D.P., "Purification and Characterization of the DNA Polymerase and RNase H Activities in Moloney Murine Sarcoma-Leukemia Virus," <i>J. Virology</i> 15(4):785-797, American Society for Microbiology (1975).

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	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>4</u>	Gerard, G.F., "Multiple RNase H Activities in Mammalian Type C Retrovirus Lysates," <i>J. Virology</i> 26(1):16-28, American Society for Microbiology (1978).
	AS	<u>4</u>	Gerard, G.F., "Mechanism of Action of Moloney Murine Leukemia Virus RNase H III," <i>J. Virology</i> 37(2):748-754, American Society for Microbiology (1981).
	AT	<u>4</u>	Gerard, G.F. et al., "Influence on Stability in <i>Escherichia coli</i> of the Carboxy-Terminal Structure of Cloned Moloney Murine Leukemia Virus Reverse Transcriptase," <i>DNA</i> 5(4):271-279, Mary Ann Liebert, Inc. (1986).

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	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

AR	<u>5</u>	Gerwin, B.I. et al., "Mutant of B-Tropic Murine Leukemia Virus Synthesizing an Altered Polymerase Molecule," <i>J. Virology</i> 31(3):741-751, American Society for Microbiology (1979).
AS	<u>5</u>	Gorecki, M. and Panet, A., "Discrimination of DNA Polymerase and RNase H Activities in Reverse Transcriptase of Avian Myeloblastosis Virus," <i>Biochemistry</i> 17(12):2438-2442, American Chemical Society (1978).
AT	<u>5</u>	Johnson, M.S. et al., "Computer analysis of retroviral pol genes: Assignment of enzymatic functions to specific sequences and homologies with nonviral enzymes," <i>Proc. Natl. Acad. Sci. USA</i> 83(20):7648-7652, National Academy of Sciences (1986).

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	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>6</u>	Kamer, G. and Argos, P., "Primary structural comparison of RNA-dependent polymerases from plant, animal and bacterial viruses," <i>Nucl. Acids. Res.</i> 12(18):7269-7282, IRL Press Ltd. (1984).
	AS	<u>6</u>	Kotewicz, M.L. et al., "Cloning and overexpression of Moloney murine leukemia virus reverse transcriptase in <i>Escherichia coli</i> ," <i>Gene</i> 35:249-258, Elsevier Science Publishers (1985).
	AT	<u>6</u>	Kotewicz, M.L. et al., "Isolation of cloned Moloney murine leukemia virus reverse transcriptase lacking ribonuclease H activity," <i>Nucl. Acids Res.</i> 16(1):265-277, IRL Press Ltd. (1988).

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	AO						Yes No
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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	7	Lai, M.-H.T. and Verma, I.M., "Reverse Transcriptase of RNA Tumor Viruses. V. In Vitro Proteolysis of Reverse Transcriptase from Avian Myeloblastosis Virus and Isolation of a Polypeptide Manifesting Only RNase H Activity," <i>J. Virology</i> 25(2):652-663, American Society for Microbiology (1978).
	AS	7	Lai, M.-H.T. et al., "Mammalian Retrovirus-Associated RNase H Is Virus Coded," <i>J. Virology</i> 27(3):823-825, American Society for Microbiology (1978).
	AT	7	Levin, J.G. et al., "Murine Leukemia Virus Mutant with a Frameshift in the Reverse Transcriptase Coding Region: Implications for pol Gene Structure," <i>J. Virology</i> 51(2):470-478, American Society for Microbiology (1984).

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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

AR	8	Maniatis, T. In: <i>Molecular Cloning, A Laboratory Manual</i> , Maniatis, T et al., eds., Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, pp. 213 and 231 (1982).
AS	8	Meloun, B. et al., "Complete primary structure of thermitase from <i>Thermoactinomyces vulgaris</i> and its structural features related to the subtilisin-type proteinases," <i>FEBS Letters</i> 183(2):195-200, North-Holland Press (1985).
AT	8	Messer, L.I. et al., "Functional Analysis of Reverse Transcription by a Frameshift pol Mutant of Murine Leukemia Virus," <i>Virology</i> 146:146-152, Academic Press, Inc. (1985).

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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	2	Michel, F. and Lang, B.F., "Mitochondrial class II introns encode proteins related to the reverse transcriptases of retroviruses," <i>Nature</i> 316:641-643, Macmillan Journals Ltd. (1985).
	AS	2	Moelling, K. and Friis, R.R., "Two Avian Sarcoma Virus Mutants with Defects in the DNA Polymerase-RNase H Complex," <i>J. Virology</i> 32(2):370-378, American Society for Microbiology (1979).
	AT	2	Resnick, R. et al., "Involvement of Retrovirus Reverse Transcriptase-Associated RNase H in the Initiation of Strong-Stop (+) DNA Synthesis and the Generation of the Long Terminal Repeat," <i>J. Virology</i> 51(3):813-821, American Society for Microbiology (1984).

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	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>10</u>	Roth, M.J. et al., "Purification and Characterization of Murine Retroviral Reverse Transcriptase Expressed in <i>Escherichia coli</i> ," <i>J. Biol. Chem.</i> 260(16):9326-9335, American Society of Biological Chemists, Inc. (1985).
	AS	<u>10</u>	Shinnick, T.M. et al., "Nucleotide sequence of Moloney murine leukaemia virus," <i>Nature</i> 293:543-548, Macmillan Journals Ltd. (1981).
	AT	<u>10</u>	Smith, J.K. et al., "Initiation of Plus-Strand DNA Synthesis During Reverse Transcription of an Avian Retrovirus Genome," <i>J. Virology</i> 49(1):200-204, American Society for Microbiology (1984).

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FORM PTO-1449 <u>INFORMATION DISCLOSURE STATEMENT</u>	ATTY. DOCKET NO. 0942.049000A/RWE/MTT	APPLICATION NO. 10/024,149
	APPLICANTS Kotewicz et al.	
	FILING DATE December 21, 2001	GROUP 1652

U.S. PATENT DOCUMENTS

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	AL						Yes No
	AM						Yes No
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	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

AR	<u>11</u>	Tanese, N. et al., "Analysis of Retroviral pol Gene Products with Antisera Raised against Fusion Proteins Produced in <i>Escherichia coli</i> ," <i>J. Virology</i> 59(2):328-340, American Society for Microbiology (1986).
AS	<u>11</u>	Tanese, N. et al., "Expression of enzymatically active reverse transcriptase in <i>Escherichia coli</i> ," <i>Proc. Natl. Acad. Sci. USA</i> 82(15):4944-4948, National Academy of Sciences (1985).
AT	<u>11</u>	Telesnitsky, A. and Goff, S.P., "RNase H domain mutations affect the interaction between Moloney murine leukemia virus reverse transcriptase and its primer-template," <i>Proc. Natl. Acad. Sci. USA</i> 90:1276-1280, National Academy of Sciences (1993).

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	AP						Yes No

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AR	<u>12</u>	Toh, H. et al., "Sequence homology between retroviral reverse transcriptase and putative polymerases of hepatitis B virus and cauliflower mosaic virus," <i>Nature</i> 305:827-829, Macmillan Journals Ltd. (1983).
AS	<u>12</u>	Tronick, S.R. et al., "Molecular cloning of Moloney murine sarcoma virus: Arrangement of virus-related sequences within the normal mouse genome," <i>Proc. Natl. Acad. Sci. USA</i> 76(12):6314-6318, National Academy of Sciences (1979).
AT	<u>12</u>	Van Beveren, C. et al., "RNA Tumor Viruses" in <i>Molecular Biology of Tumor Viruses</i> , 2nd ed., vol. 2. Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, pp. 773-779 (1985).

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	AR	<u>13</u>	Zoller, M.J. and Smith, M., "Oligonucleotide-Directed Mutagenesis of DNA Fragments Cloned into M13 Vectors," <i>Methods in Enzymology</i> 100:468-500, Academic Press, Inc. (1983).
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